

#### **PATENT**

#### IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Applicants:

Joseph L. Duffy, et al.

Serial No.:

10/701,186

Case No.: 21167

Filed:

November 4, 2003

For:

CYANTHIOPHENE DERIVATIVES, COMPOSITIONS

CONTAINING SUCH COMPOUNDS AND METHODS OF USE

Commissioner for Patents P.O. BOX 1450 Alexandria, VA 22313-1450

# INFORMATION DISCLOSURE STATEMENT UNDER 37 CFR § 1.97

Sir:

In compliance with 37 CFR § 1.97, Applicants and other individuals associated with the filing or prosecution of this application wish to bring to the attention of the Examiner the patents, publications, and other information of which they are aware, listed on the attached form PTO-1449, which they believe may be relevant to the examination of this application or may be material to patentability and in respect of which there may be a duty to disclose in accordance with 37 CFR § 1.56. Applicants respectfully request that the Examiner initial the form PTO-1449 after reviewing the pertinence of each reference.

I hereby certify that this correspondence is being deposited with the United States Postal Service as first class mail in an envelope addressed to: Commissioner for Patents, P.O. Box 1450, Alexandria, Virginia 22313-1450, on the date appearing below.

By Werck & CO., INC.

By Were por Date 2-4-2004

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This Information Disclosure Statement is not an admission that any patent, publication, or other information referred to herein is "prior art" for this invention.

In accordance with 37 CFR § 1.97(h), the filing of this Information Disclosure Statement shall not be construed to be an admission that the information cited in the Statement is, or is considered to be, material to patentability as defined in 37 CFR § 1.56(b). In accordance with 37 CFR § 1.97(g), the filing of this Information Disclosure Statement shall not be construed as a representation that a search has been made.

Pursuant to 37 CFR § 1.97(b), this Information Disclosure Statement is being filed within three months of the filing date of the present application.

Copies of the non-U.S. patent documents that are cited on the attached form PTO-1449 are enclosed herewith.

All the references cited on the attached form are in English with the exception of the following:

WO 2002/018335 A1 dated 3/7/2002 entitled "Preparation of Cyclic Amine Derivatives as CCR3 Antagonists", relates to compounds of the general formula:

$$\begin{array}{c|c}
 & R^{21} & R^{22} \\
 & N & D
\end{array}$$

JP 2000250169 A2 dated 9/14/2000 entitled Photothermographic Image Forming Material Using Diazonium Salt Coupler relates to compounds of the formula:

The material forms a coupler for a support coated with a heat-sensitive layer. It is used in thermographic imaging.

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DE 4119767 A1 dated 12/7/1992 entitled Preparation of (arenopyrimidinylthio) acetates and analogs, relates to compounds of the general formula:

In the synthesis, the compound 2-chloro-N-(3-cyano-4,5-dimethyl-2-thienyl) acetamide is disclosed.

DE 3624304 A1 dated 1/28/1988, and EP 253259A2 and A3 dated 1/20/1988 and 3/22/1989, respectively, and entitled Preparation of Substituted Thienylamides and Their Use as Agrochemical Fungicides, relate to a 3-cyanothiophene used in the synthesis schemes. The compound 3-cyano-4,5 dimethyl-2-thienyl carbamic acid methyl ester is disclosed.

DE3529247 A1 dated 11/20/1986, EP 202538 A1 dated 11/26/1986 and EP202538B1 dated 12/28/1988 and entitled Thienylureas and Isoureas and Their Preparation and Use as Growth Promoters For Animals relate to the synthesis of thienylureas and isoureas. The compound N-[3-cyano-4-(1,1-dimethylethyl)-2-thienyl]-N'-phenyl urea is disclosed.

EP193885 A1 dated 9/10/1986 and entitled Thiophene Derivatives Relates to the Preparation of Dye Intermediates. The compound N-(4-chloro-3-cyano-2-thienyl) formamide is disclosed.

EP202538 A1 dated 11/26/1986 and entitled Productivity-Increasing Agents for Livestock relates to thiophenes, thienylureas and isoureas. The compound N-[3-cyano-4-(1,1-dimethyl)-2-thienyl]-N'-phenyl urea is disclosed.

DE3507421 A1 dated 9/10/1986 and entitled Aminocyanthiophene Azo Dye Intermediates relates to azo dye intermediates of the general formula:

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The compound N-(4-chloro-3-cyano2-thienyl) formamide is disclosed.

FR2564467 A1 dated 11/22/1985 and entitled Thieno [2,3-b] Pyrroles relates to analgesics and anti-inflammatories of the general formula:

$$HO_2C-CH_2$$
 $R^3$ 
 $R^2$ 
 $R^1$ 

A cyanothiophene of the formula

is disclosed in the synthesis.

FR2565981 A1 dated 12/20/1985 and entitled Thieno [2,3-b] Pyrroles relates to thieno pyrroles of the general formula:

$$\begin{array}{c|c} R^1 & CO & R^2 \\ \hline Me & S & Me \\ \hline CHR^3R^4 & \end{array} \quad \text{and} \quad$$

A cyanthiophene of the formula:

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Thus, it is respectfully urged that the "concise explanation requirement" under 37 CFR§1.98 has been fully satisfied. If the Examiner feels that further explanation is necessary, he or she is respectfully requested to telephone the undersigned.

Any additional fees required in connection with this submission may be taken from Merck Deposit Account No. 13-2755.

Respectfully submitted,

Richard C. Billups, Reg. No. \$1,916

Attorney for Applicant

MERCK & CO., Inc.

P.O. Box 2000

Rahway, New Jersey 07065-0907

(732) 594-4683

Date: February 4, 2004

## IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Applicants:

Joseph L. Duffy, et al.

Serial No.

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For: CYANTHIOPHENE DERIVATIVES, COMPOSITIONS CONTAINING SUCH COMPOUNDS AND METHODS

OF USE

Commissioner for Patents P.O. Box 1450 Alexandria, Virginia 22313-1450

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3410 DMPOSITIONS	Examiner:	

## INFORMATION DISCLOSURE STATEMENT UNDER 37 CFR 1.97

Sir:

- 1. In compliance with 37 C.F.R. 1.97, submitted on the attached form herewith is a list of patents, publications or other information which are requested to be made of record in this application. This Information Disclosure Statement is not an admission that any patent, publication or other information referred to herein is "prior art" for this invention. In accordance with 37 C.F.R. 1.97(h), the filing of this Information Disclosure Statement shall not be construed to be an admission that the information cited in the Statement is, or is considered to be, material to patentability as defined in 37 C.F.R. 1.56(b).
- 2. In accordance with 37 C.F.R. 1.97(g), the filing of this Information Disclosure Statement shall not be construed to mean that a search has been made.
- 3. Applicants respectfully request that the Examiner initial the attached form after reviewing the pertinence of each reference.
- 4. Pursuant to the waiver by the Office of the requirement under 37 CFR 1.98 (a)(2)(i) dated July 11, 2003, if the filing date of this application is after June 30, 2003, copies of each cited U.S. patent and each U.S. patent application publication are not enclosed herewith.

I hereby certify that this correspondence is being deposited with the United States Postal Service as first class mail in an envelope addressed to: Commissioner for Patents, P.O. Box 1450, Alexandria, Virginia 22313-1450, on the date appearing below.

MERCK & CO., INC.

By Deruse brain	Date <u>2-4-2</u> 004
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### INFORMATION DISCLOSURE STATEMENT

5. Pursuant to 37 C.F.R. 1.98(d), copies of references listed on the attached form that were submitted to or cited
by the Office in a related application upon which the instant application relies for an earlier filing date under 35 U.S.C. 120
are not enclosed. Related application(s) in which references were submitted to or cited by the Office are as follows:

	RELATED APPLICATION							
U. S. SERIAL NUMBER	FILING DATE	MERCK CASE						
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If th	If this is inconvenient, additional copies will be submitted upon request.	
ı	6. In accordance with 37 C.F.R. 1.97, (check one)	
	the attached information is filed within three months of the filing date of the captioned case.	
	the attached information is filed more than three months after the filing date but prior to the mailing of Office Action on the merits.	a first
	the attached information is being filed more than three months after the filing date and after the mailing.  Office Action on the merits, but before the mailing date of a Final Action or Notice of Allowance. The authorization is therefore given to charge Deposit Account No. 13-2755 for the fee required under 37	enclosed
	1.17(p). the undersigned certifies that each item of information contained in this Information Disclosure Statem cited in any communication from a foreign patent office in a counterpart foreign application not more to months prior to the filing of this Statement.	nent was first nan three
	the undersigned certifies that no item of information contained in this Information Disclosure Statemer a communication from a foreign patent office in a counterpart foreign application, and, to the knowled person signing the certification after making reasonable inquiry, was known to any individual designate C.F.R. 1.56(c) more than three months prior to the filing of this Statement.	ge of the
	Respectfully submitted,  By: Richard C. Billups  Attorney For Applicant(s)  Reg. No. 31,916  MERCK & CO., INC.  Patent Dept., RY60-30  P.O. Box 2000  Rahway, N.J. 07065-0907  (732)594-4683	7

Date: February 4, 2004

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- 4	Substitute	for form 1449Å/PTO	•		COMPLETE IF KNOWN				
	DANKODEW	PRMATION	DIS	CLOSURE	Application Number	10/701,186			
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	STATEMENT BY APPLICANT				First Named Inventor	Joseph L. Duffy et al.			
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*	Sheet	1	of	5	Attorney Docket Number	21167			

U.S. PATENT DOCUMENTS										
Cited Docui	ıment	Date of Publicati Cited Docume MM-DD-YYY	Applicant nent	Name of Patentee or Ap of Cited Documer	Kind Code	Document I	U.S. Patent I Number	Cite No.	Examiner Initials*	
7/1998		07/07/1998		Stephen E. de Laszlo, et al.			5,776,954	1		
6/1982		11/16/1982		Paul L. Anderson, et al.			4,359,474	2		
8/1986		01/28/1986		Michel Wierzbicki	ļ		4,567,196	3		
7/1976		04/27/1976		William J. Wechter, et al.			3,953,468	4		
6/1986		08/26/1986		Michael Wierzbicki, et al.			4,608,384	5		
6/1989		05/16/1989		John E. Francis			4,831,013	6	•	
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	FOREIGN PATENT DOCUMENTS										
Examiner Cite			Foreign Patent Document		Name of Patentee or Applicant	Date of Publication of					
Initials*	No.	Office		Kind Code (if known)	of Cited Document	Cited Document MM-DD-YYYY					
	7	PCT	WO 00/69810		NOVO NORDISK A/S & AGOURON PHARMACEUTICALS, INC.	11/23/2000					
	8	PCT	WO 00/39088		NOVO NORDISK A/S & AGOURON PHARMACEUTICALS, INC.	07/06/2000					
	9	PCT	WO 99/01423corrected version		NOVO NORDISK A/S & ALANEX CORPORATION	01/14/1999					
	10	PCT	WO 99/01423		NOVO NORDISK A/S & ALANEX CORPORATION	01/14/1999					
	11	PCT	WO 98/04528 A2		BAYER CORPORATION & BAYER AKTIENGESELLSCHAFT	02/05/1998					
	12	PCT	WO 98/04528 A3		BAYER CORPORATION & BAYER AKTIENGESELLSCHAFT	02/05/1998					
	13	PCT	WO 99/22108		MERCK & CO., INC.	05/28/1998					
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	15	РСТ	WO 97/16442		MERCK & CO., INC.	05/09/1997					
	16	PCT	WO 98/21957		MERCK & CO., INC.	05/28/1998					

Examiner Signature Date Considered

<sup>\*</sup>Examiner: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.

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	LOEWE	VFORMATION I	DIS	CLOSURE	Application Number	10/701,186	
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		TAZEMENT DI	AF	PLICANI	First Named Inventor	Joseph L. Duffy, et al.	
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U.S. PATENT DOCUMENTS								
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	FOREIGN PATENT DOCUMENTS									
Examiner	Cite		Foreign Patent Document		Name of Patentee or Applicant	Date of Publication of				
Initials*	No.	Office		Kind Code if known)	of Cited Document	Cited Document MM-DD-YYYY				
	17	PCT	WO 99/46267		NOVO NORDISK A/S	09/16/1999				
	18	PCT	WO 99/32448		AMGEN, INC.	07/01/1999				
	19	PCT	WO 99/32448 corrected version		AMGEN, INC.	07/01/1998				
	20	PCT	WO 02/18335		YAMANOUCHI PHARMACEUTICAL CO., LTD.	03/07/2002				
	21	PCT	WO 01/58890		ASTRAZENECA AB	08/16/2001				
	22	PCT	WO 01/46165		NOVARTIS-ERFINDUNGEN VERWALTUNGSGESELLSCHAFT M.B.H.	06/28/2001				
	23	PCT	WO 01/44226		PROTHERICS MOLECULAR DESIGN LIMITED	06/21/2001				
	24	PCT	WO 01/40223 A2		NOVARTIS AG	06/07/2001				
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	U.S. PATENT DOCUMENTS							
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			FOREI	GN PATENT	T DOCUMENTS	
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	27	PCT	WO 01/32632 A2	(if known)	ELI LILLY AND COMPANY	05/10/2001
	28	PCT	WO 01/32632 A3		ELI LILLY AND COMPANY	05/10/2001
	29	EP	0 263 071 A1		CIBA-GEIGY AG	04/06/1988
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	INFO	<b>DRMATION</b>	DISC	CLOSURE	Application Number	10/701,186		
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3	STATEMENT BY APPLICANT			FLICANI	First Named Inventor	Joseph L. Duffy, et al.		
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U.S. PATENT DOCUMENTS						
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	45	JP	2002-50169		FUJI PHOTO FILM CO., LTD.	02/15/2002			

Examiner Signature	Date Considered	

<sup>\*</sup>Examiner: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.

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COMPLETE IF KNOWN Substitute for form 1449B/PTO FORMATION DISCLOSURE 10/701,186 Application Number Filing Date November 4, 2003 **TEMENT BY APPLICANT** First Named Inventor Joseph L. Duffy et al. Group Art Unit (use as many sheets as necessary) Examiner Name 5 5 21167 of Attorney Docket Number

Cite No.	Include name of the author, title, date, page(s), volume-issue number(s) and place of publication.
16	Guillon, Jean, et al., "Synthesis of new pyrrolo[1,2-a]quinoxalines: potential non-peptide glucagon receptor antagonist", Eur. J. Med. Chem, 33, pp. 293-308 (1998)
17	Petersen, K. F., et al., "Effects of a novel glucagon receptor antagonist (Bay 27-9955) on glucagon-stimulated glucose production in humans", <i>Diabetologia</i> , 44, pp. 2018-2024 (2001)
18	Ladouceur, Gaetan, H., et al., "Discovery of 5-Hydroxyalkyl-4-phenylpyridines as a New Class of Glucagon Receptor Antagonists", <i>Bioorganic &amp; Medicinal Chemistry Letters</i> , 12, pp. 461-464 (2002)
19	Cascieri, Margaret, A., "Characterization of a Novel, Non-peptidyl Antagonist of the Human Glucagon Receptor, <i>The Journal of Biological Chemistry</i> , Vol. 274, Issue of March 26, pp. 8694-8697 (1999)
50	de Laszlo, Stephen, E., "Potent, Orally Absorbed Glucagon Receptor Antagonists", <i>Bioorganic &amp; Medicinal Chemistry Letters</i> , 9, pp. 641-646 (1999)
51	Madsen, Peter, et al., "Discovery and Structure-Activity Relationship of the First Non-Peptide Competitive Human glucagon Receptor Antagonists", <i>J. Med Chem</i> , 41, pp. 5150-5157 (1998)
52	Ling, Anthony, et al., "Identification of Alkylidene Hydrazides as Glucagon Receptor Antagonists", <i>J. Med. Chem</i> , 44, pp. 3141-3149 (2001)
53	Ling, Anthony, et al., "Human Glucagon Receptor Antagonists Based on Alkylidine Hydrazides", <i>Bioorganic &amp; Medicinal Chemistry Letters</i> , 12, pp. 663-666 (2002)
54	Collins, Judith, L., "CP-99,711: A Non-Peptide Glucagon Receptor Antagonist", Bioorganic & Medicinal Chemistry Letters, Vol. 2, No. 9, pp. 915-918 (1992)
55	Parker, Janice, C., "Effects of Skyrin, a Receptor-Selective Glucagon Antagonist, in Rat and human Hepatocytes", <i>Diabetes</i> , Vol. 49, pp. 2079-2086 (December 2000)
56	Connell, Richard, D., "Glucagon antagonists for the treatment of Type 2 diabetes", Expert Opinion on Therapeutic patents, 9(6), pp. 701-709 (1999)
57	Madsen, P., et al., "Advances in Non-Peptide Glucagon Receptor Antagonists", Current Pharmaceutical Design, 5, pp. 683-691 (1999)
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